PATENT Docker No.: SHIM1120

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:

AIZAWA et al

Group No.: Unknown

Serial No.:

09/830,019

Examiner: Unknown

Filed:

April 19, 2001

For: VACCINE PREPARATIONS CONTAINING ATTENUATED TOXIN

BOX PCT

Commissioner for Patents Washington, D.C. 20231

Sir:

PRELIMINARY AMENDMENT

In advance of the first examination in this case, Applicants request amendment of the subject application, as follows:

"EXPRESS MAIL" Mailing Label Number EL617043151US

Date of Deposit August 7, 2001_

I hereby certify under 37 CFR 1.10 that this correspondence is being deposited with the United States Postal Service as "Express Mail Post Office to Addressee" with sufficient postage on the date indicated above and is addressed to the Commissioner for Patrils, Washington,

D.C. 20231

Tim Ellis

IN THE SPECIFICATION

Please amend the specification by inserting the following replacement paragraphs:

Please replace the paragraph at page 16, lines 18 to 34 with the following paragraph:

--Because a mutant toxin has a partially altered structure as compared with that of the natural molecule, it generally loses some of the original properties of the toxin as a result of such structural changes. However, there are cases in which the activity of enhancing immunity is retained. When the toxic activity of mutant toxin is sufficiently low, e.g., reduced by a factor of at least one-two thousandth relative to that of the natural one, it can be used as an attenuated toxin for the adjuvant of the present invention provided its activity of enhancing immunity is verified. Otherwise, the mutant toxin is further attenuated by using any of the above-mentioned chemical and physical treatments to give rise to a usable

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